#### WHAT IS CLAIMED IS:

### 1. A compound of structural formula I:

$$R^{5}O$$
 $X$ 
 $B$ 
 $R^{4}$ 
 $X$ 
 $R^{6}$ 
 $R^{4}$ 
 $R^{3}$ 
 $R^{2}$ 
(I)

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or a pharmaceutically acceptable salt thereof; wherein n is 0, 1, or 2;
B is

10 X is CH2, CHF, CF2, or C=CH2;

Y is N or C-R<sup>9</sup>;

W is O or S;

 $R^1$  is  $C_{2-4}$  alkenyl,  $C_{2-4}$  alkynyl, or  $C_{1-4}$  alkyl, wherein alkyl is unsubstituted or substituted with hydroxy, amino,  $C_{1-4}$  alkoxy,  $C_{1-4}$  alkylthio, or one to three fluorine

15 atoms;

 $R^2$  is hydrogen, fluorine, amino, hydroxy, mercapto,  $C_{1-4}$  alkoxy,  $C_{1-8}$  alkylcarbonyloxy, or  $C_{1-4}$  alkyl;

R<sup>3</sup> and R<sup>4</sup> are each independently selected from the group consisting of hydrogen, cyano, azido, halogen, hydroxy, mercapto, amino, C<sub>1-4</sub> alkoxy, C<sub>1-8</sub>

alkylcarbonyloxy, C<sub>2-4</sub> alkenyl, C<sub>2-4</sub> alkynyl, and C<sub>1-4</sub> alkyl, wherein alkyl is unsubstituted or substituted with hydroxy, amino, C<sub>1-4</sub> alkoxy, C<sub>1-4</sub> alkylthio, or one to three fluorine atoms;

R5 is hydrogen, C1-10 alkylcarbonyl, P3O9H4, P2O6H3, or P(O)R13R14;

R<sup>6</sup> and R<sup>7</sup> are each independently hydrogen, methyl, hydroxymethyl, or fluoromethyl; R<sup>8</sup> is hydrogen, C<sub>1-4</sub> alkyl, C<sub>2-4</sub> alkynyl, halogen, cyano, carboxy, C<sub>1-4</sub> alkyloxycarbonyl, azido, amino, C<sub>1-4</sub> alkylamino, di(C<sub>1-4</sub> alkyl)amino, hydroxy, C<sub>1-6</sub> alkoxy, C<sub>1-6</sub> alkylthio, C<sub>1-6</sub> alkylsulfonyl, or (C<sub>1-4</sub> alkyl)<sub>0-2</sub> aminomethyl; R<sup>9</sup> is hydrogen, halogen, cyano, nitro, NHCONH<sub>2</sub>, CONR<sup>12</sup>R<sup>12</sup>, CSNR<sup>12</sup>R<sup>12</sup>, COOR<sup>12</sup>, C(=NH)NH<sub>2</sub>, hydroxy, C<sub>1-3</sub> alkoxy, amino, C<sub>1-4</sub> alkylamino, di(C<sub>1-4</sub> alkyl)amino, or C<sub>1-3</sub> alkyl, wherein alkyl is unsubstituted or substituted with one to three groups independently selected from halogen, amino, hydroxy, carboxy, and C<sub>1-3</sub> alkoxy;

10 R<sup>10</sup> and R<sup>16</sup> are each independently hydrogen, hydroxy, mercapto, halogen, C<sub>1-4</sub> alkoxy, C<sub>1-4</sub> alkylthio, C<sub>1-8</sub> alkylcarbonyloxy, C<sub>3-6</sub> cycloalkylcarbonyloxy, C<sub>1-8</sub> alkyloxycarbonyloxy, C<sub>3-6</sub> cycloalkyloxycarbonyloxy, -OCH<sub>2</sub>CH<sub>2</sub>SC(=O)C<sub>1-4</sub> alkyl, -OCH<sub>2</sub>O(C=O)C<sub>1-4</sub> alkyl, -OCH(C<sub>1-4</sub> alkyl)O(C=O)C<sub>1-4</sub> alkyl, amino, C<sub>1-4</sub> alkylamino, di(C<sub>1-4</sub> alkyl)amino, C<sub>3-6</sub> cycloalkylamino, di(C<sub>3-6</sub> cycloalkyl)amino,

or an amino acyl residue having structural formula

$$R^{20}$$
 O  $R^{17}$  or  $R^{20}$  O  $R^{18}R^{19}$ 

R<sup>11</sup> is hydrogen, hydroxy, halogen, C<sub>1-4</sub> alkoxy, amino, C<sub>1-4</sub> alkylamino, di(C<sub>1-4</sub> alkyl)amino, C<sub>3-6</sub> cycloalkylamino, or di(C<sub>3-6</sub> cycloalkylamino); each R<sup>12</sup> is independently hydrogen or C<sub>1-6</sub> alkyl;

20 R<sup>17</sup>, R<sup>18</sup>, and R<sup>19</sup> are each independently hydrogen or C<sub>1-6</sub> alkyl; R<sup>13</sup> and R<sup>14</sup> are each independently hydroxy, -OCH<sub>2</sub>CH<sub>2</sub>SC(=O)C<sub>1-4</sub> alkyl, -OCH<sub>2</sub>O(C=O)OC<sub>1-4</sub> alkyl, -NHCHMeCO<sub>2</sub>Me, -OCH(C<sub>1-4</sub> alkyl)O(C=O)C<sub>1-4</sub> alkyl,

R<sup>15</sup> is hydrogen, C<sub>1-6</sub> alkyl, C<sub>2-6</sub> alkenyl, C<sub>2-6</sub> alkynyl, C<sub>1-4</sub> alkylamino, CF<sub>3</sub>, or halogen; and R<sup>20</sup> is hydrogen, C<sub>1-4</sub> alkyl, or phenyl C<sub>0-2</sub> alkyl; with the proviso that when B is

X is CH2; Y is N;  $R^{10}$  is NH2;  $R^2$  and  $R^3$  are  $\alpha$ -OH; and  $R^4$ ,  $R^5$ ,  $R^6$ ,  $R^7$ ,  $R^8$ , and  $R^{11}$  are hydrogen, then  $R^1$  is not  $\beta$ -methyl.

#### 2. The compound of Claim 1 wherein B is

$$R^8 \longrightarrow N \longrightarrow N \longrightarrow R^{10}$$

#### 3. The compound of Claim 2 of structural formula II:

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 $R^1$  is  $C_{1-3}$  alkyl, wherein alkyl is unsubstituted or substituted with one to three fluorine atoms;

 $R^2$  is hydroxy, fluoro,  $C_{1-3}$  alkoxy, or  $C_{1-8}$  alkylcarbonyloxy;

R<sup>3</sup> is hydrogen, halogen, hydroxy, amino, C<sub>1-3</sub> alkoxy, or C<sub>1-8</sub> alkylcarbonyloxy;

15 R<sup>5</sup> is hydrogen, C<sub>1-8</sub> alkylcarbonyl, P<sub>3</sub>O<sub>9</sub>H<sub>4</sub>, P<sub>2</sub>O<sub>6</sub>H<sub>3</sub>, or PO<sub>3</sub>H<sub>2</sub>;

R8 is hydrogen, amino, or C1-4 alkylamino; and

 $R^{10}$  and  $R^{11}$  are each independently hydrogen, halogen, hydroxy, amino,  $C_{1-4}$  alkylamino, di( $C_{1-4}$  alkylamino, or  $C_{3-6}$  cycloalkylamino; with the proviso that when  $R^{10}$  is NH<sub>2</sub>,  $R^2$  and  $R^3$  are  $\alpha$ -OH, and  $R^5$ ,  $R^8$ , and  $R^{11}$  are hydrogen, then  $R^1$  is not  $\beta$ -methyl.

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#### 4. The compound of Claim 3 wherein

R1 is methyl, fluoromethyl, difluoromethyl, or trifluoromethyl;

R<sup>2</sup> is hydroxy, fluoro, or methoxy;

R<sup>3</sup> is hydrogen, fluoro, hydroxy, amino, or methoxy;

10 R<sup>5</sup> is hydrogen or P<sub>3</sub>O<sub>9</sub>H<sub>4</sub>;

R8 is hydrogen or amino; and

 $R^{10}$  and  $R^{11}$  are each independently hydrogen, fluoro, hydroxy, or amino; with the proviso that when  $R^{10}$  is NH<sub>2</sub>,  $R^2$  and  $R^3$  are  $\alpha$ -OH, and  $R^5$ ,  $R^8$ , and  $R^{11}$  are hydrogen, then  $R^1$  is not  $\beta$ -methyl.

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### 5. The compound of Claim 2 of structural formula III:

wherein

R<sup>1</sup> is C<sub>1-3</sub> alkyl, wherein alkyl is unsubstituted or substituted with one to three

20 fluorine atoms;

R<sup>2</sup> is hydroxy, fluoro, C<sub>1-3</sub> alkoxy, or C<sub>1-8</sub> alkylcarbonyloxy;

R<sup>3</sup> is hydrogen, halogen, hydroxy, amino, C<sub>1-3</sub> alkoxy, or C<sub>1-8</sub> alkylcarbonyloxy;

R<sup>5</sup> is hydrogen, C<sub>1-8</sub> alkylcarbonyl, P<sub>3</sub>O<sub>9</sub>H<sub>4</sub>, P<sub>2</sub>O<sub>6</sub>H<sub>3</sub>, or PO<sub>3</sub>H<sub>2</sub>;

R8 is hydrogen, amino, or C1-4 alkylamino;

R<sup>9</sup> is hydrogen, cyano, methyl, halogen, CONH<sub>2</sub> or CSNH<sub>2</sub>; and R<sup>10</sup> and R<sup>11</sup> are each independently hydrogen, halogen, hydroxy, amino,

C<sub>1-4</sub> alkylamino, di(C<sub>1-4</sub> alkyl)amino, or C<sub>3-6</sub> cycloalkylamino.

6. The compound of Claim 5 wherein

R1 is methyl, fluoromethyl, difluoromethyl, or trifluoromethyl;

 $R^2$  is hydroxy, fluoro, or methoxy;

R3 is hydrogen, fluoro, hydroxy, amino, or methoxy;

R5 is hydrogen or P3O9H4;

R8 is hydrogen or amino;

R9 is hydrogen, cyano, methyl, halogen, CONH2 or CSNH2; and

10 R10 and R11 are each independently hydrogen, fluoro, hydroxy, or amino.

# 7. The compound of Claim 2 of structural formula IV:

$$R^{5}O$$
 $R^{8}$ 
 $N$ 
 $N$ 
 $N$ 
 $N$ 
 $R^{11}$ 
 $R^{10}$ 
 $R^{11}$ 
 $R^{10}$ 
 $R^{11}$ 
 $R^{11}$ 

wherein

R<sup>1</sup> is C<sub>1-3</sub> alkyl, wherein alkyl is unsubstituted or substituted with one to three fluorine atoms;

R2 is hydroxy, fluoro, C1-3 alkoxy, or C1-8 alkylcarbonyloxy;

R<sup>3</sup> is hydrogen, halogen, hydroxy, amino, C<sub>1-3</sub> alkoxy, or C<sub>1-8</sub> alkylcarbonyloxy;

R5 is hydrogen, C1-8 alkylcarbonyl, P3O9H4, P2O6H3, or PO3H2;

20 R8 is hydrogen, amino, or C1-4 alkylamino; and

R10 and R11 are each independently hydrogen, halogen, hydroxy, amino,

C<sub>1-4</sub> alkylamino, di(C<sub>1-4</sub> alkyl)amino, or C<sub>3-6</sub> cycloalkylamino.

8. The compound of Claim 7 wherein

25 R<sup>1</sup> is methyl, fluoromethyl, difluoromethyl, or trifluoromethyl; R<sup>2</sup> is hydroxy, fluoro, or methoxy;

R3 is hydrogen, fluoro, hydroxy, amino, or methoxy;

R<sup>5</sup> is hydrogen or P<sub>3</sub>O<sub>9</sub>H<sub>4</sub>;

R8 is hydrogen or amino; and

R10 and R11 are each independently hydrogen, fluoro, hydroxy, or amino.

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## 9. The compound of Claim 2 of structural formula V:

$$R^{5}O$$
 $R^{8}$ 
 $R^{9}$ 
 $R^{10}$ 
 $R^{10}$ 
 $R^{11}$ 
 $R^{11}$ 

wherein

R1 is C1-3 alkyl, wherein alkyl is unsubstituted or substituted with one to three

10 fluorine atoms;

R2 is hydroxy, fluoro, C1-3 alkoxy, or C1-8 alkylcarbonyloxy;

R3 is hydrogen, halogen, hydroxy, amino, C1-3 alkoxy, or C1-8 alkylcarbonyloxy;

R5 is hydrogen, C<sub>1-8</sub> alkylcarbonyl, P<sub>3</sub>O<sub>9</sub>H<sub>4</sub>, P<sub>2</sub>O<sub>6</sub>H<sub>3</sub>, or PO<sub>3</sub>H<sub>2</sub>;

R8 is hydrogen, amino, or C1-4 alkylamino;

15 R9 is hydrogen, cyano, methyl, halogen, CONH2 or CSNH2; and

R10 and R11 are each independently hydrogen, halogen, hydroxy, amino,

C<sub>1-4</sub> alkylamino, di(C<sub>1-4</sub> alkyl)amino, or C<sub>3-6</sub> cycloalkylamino.

## 10. The compound of Claim 9 wherein

20 R1 is methyl, fluoromethyl, difluoromethyl, or trifluoromethyl;

R<sup>2</sup> is hydroxy, fluoro, or methoxy;

R<sup>3</sup> is hydrogen, fluoro, hydroxy, amino, or methoxy;

R<sup>5</sup> is hydrogen or P<sub>3</sub>O<sub>9</sub>H<sub>4</sub>;

R8 is hydrogen or amino;

R9 is hydrogen, cyano, methyl, halogen, CONH2 or CSNH2; and R10 and R11 are each independently hydrogen, fluoro, hydroxy, or amino.

11. The compound of Claim 2 selected from the group consisting of:

- 5 2-amino-7-[ $(1\beta,2\alpha OH,3\alpha,4\beta)$ -2,3-dihydroxy-4-hydroxymethyl-2-methyl-5-methylenecyclopentyl]-3,7-dihydro-4*H*-pyrrolo[2,3-d]pyrimidin-4-one;
  - 2-amino-7-[(1R,2S,3R,4R)-2,3-dihydroxy-4-hydroxymethyl-2-methyl-5-methylenecyclopentyl]-3,7-dihydro-4*H*-pyrrolo[2,3-d]pyrimidin-4-one;

10 (1αOH,2α,3β,5β)-5-(4-amino-7*H*-pyrrolo[2,3-d]pyrimidin-7-yl)-3-hydroxymethyl-1-methyl-4- methylenecyclopentane-1,2-diol;

- (1S,2R,3R,5R)-5-(4-amino-7*H*-pyrrolo[2,3-d]pyrimidin-7-yl)-3-hydroxymethyl-1methyl-4- methylenecyclopentane-1,2-diol;
  - $(1\beta,2\alpha OH,3\alpha,4\beta)$ -2-amino-9-[2,3-dihydroxy-4-(hydroxymethyl)-2-methyl-5-methylenecyclopentyl]-1,9-dihydro-6H-purin-6-one;
- 20 2-amino-9-[(1R,2S,3R,4R)-2,3-dihydroxy-4-(hydroxymethyl)-2-methyl-5-methylenecyclopentyl]-1,9-dihydro-6*H*-purin-6-one;

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- (1S,2R,3R,5R)-5-(6-amino-9*H*-purin-9-yl)-3-(hydroxymethyl)-1-methyl-4-methylenecyclopentane-1,2-diol;
- $(1\alpha OH, 2\alpha, 3\beta, 5\beta)$ -5-(6-amino-9H-purin-9-yl)-3-(hydroxymethyl)-1-methyl-4-methylenecyclopentane-1,2-diol;
- (1RS,2R,3R,5R)-5-(4-amino-7*H*-pyrrolo[2,3-*d*]pyrimidin-7-yl)-3-(hydroxymethyl)-1-30 methylcyclopentanediol-1,2-diol;
  - (1S,2R,3R,5R)-5-(4-amino-7*H*-pyrrolo[2,3-*d*]pyrimidin-7-yl)-3-(hydroxymethyl)-1-methylcyclopentanediol-1,2-diol;

(1RS,2R,3R,5R)-5-(6-amino-9*H*-purin-9-yl)-3-(hydroxymethyl)-1-methylcyclopentanediol-1,2-diol;

- (1S,2R,3R,5R)-5-(6-amino-9*H*-purin-9-yl)-3-(hydroxymethyl)-1-5 methylcyclopentanediol-1,2-diol;
  - 2-amino-9-[(1R,2RS,3R,4R)-2,3-dihydroxy-4-(hydroxymethyl)-2-methylcyclopentyl]-1,9-dihydro-6*H*-purin-6-one;
- 2-amino-9-[(1R,2S,3R,4R)-2,3-dihydroxy-4-(hydroxymethyl)-2-methylcyclopentyl]-1,9-dihydro-6*H*-purin-6-one;
  - 2-amino-7-[(1R,2RS,3R,4R)-2,3-dihydroxy-4-(hydroxymethyl)-2-methylcyclopentyl]-3,7-dihydro-4*H*-pyrrolo[2,3-*d*]pyrimidin-4-one; and

2-amino-7-[(1R,2S,3R,4R)-2,3-dihydroxy-4-(hydroxymethyl)-2-methylcyclopentyl]-3,7-dihydro-4*H*-pyrrolo[2,3-*d*]pyrimidin-4-one;

and the corresponding 5'-triphosphates; or a pharmaceutically acceptable salt thereof

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- 12. A pharmaceutical composition comprising a compound of Claim 1 and a pharmaceutically acceptable carrier.
- 25 13. A method of treating RNA-dependent RNA virus infection comprising administering to a mammal in need of such treatment a therapeutically effective amount of a compound according to Claim 1.
- 14. The method of Claim 13 wherein said RNA-dependent RNA virus infection is a hepatitis C virus (HCV) infection.
  - 15. The method of Claim 14 in combination with a therapeutically effective amount of another agent active against HCV.

16. The method of Claim 15 wherein said agent active against HCV is a 2'-C-Me-ribonucleoside; ribavirin; levovirin; thymosin alpha-1; interferon- $\beta$ ; an inhibitor of NS3 serine protease; an inhibitor of inosine monophosphate dehydrogenase; interferon- $\alpha$  or pegylated interferon- $\alpha$ , alone or in combination with ribavirin or levovirin.

- 17. The method of Claim 16 wherein said agent active against HCV is interferon-α or pegylated interferon-α, alone or in combination with ribavirin.
- 18. Use of a compound of Claim 1 for treatment of RNA-dependent RNA virus infection in a mammal.

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- 19. The use of Claim 18 wherein said RNA-dependent RNA virus infection is HCV infection.
- 20. Use of a compound of Claim 1 in the manufacture of a medicament for treatment of RNA-dependent RNA virus infection in a mammal.
- 21. The use of Claim 20 wherein said RNA-dependent RNA virus 20 infection is HCV infection.